Absolute stereochemistry.

$$Me_3+N$$
 $NH_2$ 
 $R$ 
 $NH_2$ 
 $S$ 
 $CO_2H$ 

RN 853015-50-6 CAPLUS

Absolute stereochemistry.

=> d his

L1

(FILE 'HOME' ENTERED AT 17:46:50 ON 15 SEP 2007)

FILE 'REGISTRY' ENTERED AT 17:47:02 ON 15 SEP 2007

STRUCTURE UPLOADED

L2 0 S L1 SSS SAM

L3 6 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 17:48:42 ON 15 SEP 2007

L4 4 S L3

=> d his

(FILE 'HOME' ENTERED AT 17:46:50 ON 15 SEP 2007)

FILE 'REGISTRY' ENTERED AT 17:47:02 ON 15 SEP 2007

L1 STRUCTURE UPLOADED

L2 ----- 0 S L-1 -- SSS -- SAM ----

L3 6 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 17:48:42 ON 15 SEP 2007

L4 4 S L3

=> d his

(FILE 'HOME' ENTERED AT 17:46:50 ON 15 SEP 2007)

FILE 'REGISTRY' ENTERED AT 17:47:02 ON 15 SEP 2007

L1 STRUCTURE UPLOADED

L2 0 S L1 SSS SAM

L3 6 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 17:48:42 ON 15 SEP 2007

Copylor JULIO, POS

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Welcome to STN International! Enter x:x
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         MAY 14
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         MAY 21
                 CA/CAplus enhanced with additional kind codes for German
                 patents
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         MAY 22
                 CA/CAplus enhanced with IPC reclassification in Japanese
                 patents
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         JUN 27
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        JUL 02
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NEWS 14 JUL 02
                 SCISEARCH enhanced with complete author names
NEWS 15 JUL 02
                 CHEMCATS accession numbers revised
NEWS 16 JUL 02
                 CA/CAplus enhanced with utility model patents from China
NEWS 17
        JUL 16
                 CAplus enhanced with French and German abstracts
NEWS 18 JUL 18
                 CA/CAplus patent coverage enhanced
NEWS 19 JUL 26
                 USPATFULL/USPAT2 enhanced with IPC reclassification
NEWS 20 JUL 30
                 USGENE now available on STN
NEWS 21 AUG 06
                 CAS REGISTRY enhanced with new experimental property tags
NEWS 22 AUG 06
                 BEILSTEIN updated with new compounds
NEWS 23
        AUG 06
                 FSTA enhanced with new thesaurus edition
NEWS 24 AUG 13
                 CA/CAplus enhanced with additional kind codes for granted
                 patents
         AUG 20
NEWS 25
                 CA/CAplus enhanced with CAS indexing in pre-1907 records
NEWS 26
        AUG 27
                 Full-text patent databases enhanced with predefined
                 patent family display formats from INPADOCDB
NEWS 27
         AUG 27
                 USPATOLD now available on STN
NEWS -28 AUG 28
                 CAS REGISTRY enhanced with additional experimental
                 spectral property data
NEWS 29
                 STN AnaVist, Version 2.0, now available with Derwent
         SEP 07
                 World Patents Index
NEWS 30
        SEP 13
                 FORIS renamed to SOFIS
NEWS 31
        SEP 13
                 INPADOCDB: New SDI frequency MONTHLY available now
             05 SEPTEMBER 2007: CURRENT WINDOWS VERSION IS V8.2,
NEWS EXPRESS
              CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
              AND CURRENT DISCOVER FILE IS DATED 05 SEPTEMBER 2007.
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=> Uploading A:\10.580803.Sheu et al..str

L1 STRUCTURE UPLOADED

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L1 HAS NO ANSWERS

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G1 [@1], [@2]

Structure attributes must be viewed using STN Express query preparation.

=> s 11 sss sam

SAMPLE SEARCH INITIATED 17:47:41 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 15 TO ITERATE

100.0% PROCESSED

15 ITERATIONS

0 ANSWERS

6 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 68 TO 532

PROJECTED ANSWERS: 0 TO (

L2 0 SEA SSS SAM L1

=>-s-l1-sss-full

FULL SEARCH INITIATED 17:47:50 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 245 TO ITERATE

100.0% PROCESSED 245 ITERATIONS SEARCH TIME: 00.00.01

L3 6 SEA SSS FUL L1

=> d scan

L3 6 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN Ethanaminium, 2-[(2R)-2-(acetylamino)-3-mercapto-1-oxopropoxy]-N,N,N-trimethyl-, chloride (1:1)

MF C10 H21 N2 O3 S . C1

Absolute stereochemistry.

• cl -

## \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):end

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COST IN U.S. DOLLARS

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FULL ESTIMATED COST

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(FILE 'HOME' ENTERED AT 17:46:50 ON 15 SEP 2007)

FILE 'REGISTRY' ENTERED AT 17:47:02 ON 15 SEP 2007

L1 STRUCTURE UPLOADED

L2 0 S L1 SSS SAM

L3 6 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 17:48:42 ON 15 SEP 2007

=> s 13

L4 4 L3

=> d l4 ed ibib abs hitstr 1-4

L4 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

ED Entered STN: 20 May 2007

ACCESSION NUMBER: 2007:542355 CAPLUS

DOCUMENT NUMBER: 147:157119

TITLE: Targeting antioxidants to mitochondria: a potential

new therapeutic strategy for cardiovascular diseases

AUTHOR(S): Victor, V. M.; Rocha, M.

CORPORATE SOURCE: Centro Nacional de Investigaciones Cardiovasculares

(CNIC), Madrid, 28029, Spain

SOURCE: Current Pharmaceutical Design (2007), 13(8), 845-863

CODEN: CPDEFP; ISSN: 1381-6128 Bentham Science Publishers Ltd.

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

PUBLISHER:

A review. Mitochondria produce large amts. of free radicals and play an important role in the life and death of a cell. Thus, mitochondrial oxidative damage and dysfunction contribute to a number of cell pathologies that manifest themselves through a range of conditions including ischemia-reperfusion injury, sepsis, diabetes, atherosclerosis and, consequently, cardiovascular diseases (CVD). In fact, endothelial dysfunction, characterized by a loss of nitric oxide (NO) bioactivity, occurs early on in the development of atherosclerosis, and dets. future vascular complications. Although the mol. mechanisms responsible for mitochondria-mediated disease processes are not yet clear, oxidative stress seems to play an important role. This review considers the process of CVD from a mitochondrial perspective. Accordingly, strategies for the targeted delivery of antioxidants to mitochondria are being developed. In this review, we will provide a summary of the following areas: the cellular metabolism of reactive oxygen species (ROS) and its role in pathophysiol. processes such as CVD; currently available antioxidants and possible reasons for their efficacy and inefficacy in ameliorating oxidative stress-mediated diseases; recent developments in mitochondrially-targeted antioxidants that concentrate on the matrix-facing surface of the inner mitochondrial membrane and therefore protect against mitochondrial oxidative damage, and their therapeutic potential for future treatment of CVDs. More pre-clin. and clin. studies, however, are necessary in order to evaluate the effectiveness and toxicity of mitochondrially-targeted antioxidants.

IT 853015-46-0, MitoNAC 943963-94-8, MitoGSH

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(targeting antioxidants to mitochondria with a potential new therapeutic strategy for cardiovascular diseases)

RN 853015-46-0 CAPLUS

CN Ethanaminium, 2-[(2R)-2-(acetylamino)-3-mercapto-1-oxopropoxy]-N,N,N-trimethyl-, chloride (1:1) (CA INDEX NAME)

Absolute stereochemistry.

● Cl -

RN 943963-94-8 CAPLUS CN INDEX NAME NOT YET ASSIGNED

## Absolute stereochemistry.

REFERENCE COUNT:

156 THERE ARE 156 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE **FORMAT** 

ANSWER 2 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN L4

USA

ED Entered STN: 21 Jul 2006

ACCESSION NUMBER: 2006:710737 - CAPLUS

DOCUMENT NUMBER:

145:146030

TITLE:

Preparation of compounds for delivering amino acids or peptides with antioxidant activity into mitochondria

INVENTOR (S):

Sheu, Shey-Shing; Anders, Marion W.; Xu, Lin; Sharma,

Virendra K.; Nauduri, Dhananjaya

PATENT ASSIGNEE(S):

SOURCE:

U.S. Pat. Appl. Publ., 15 pp., Cont.-in-part of Appl.

No. PCT/US04/039739.

CODEN: USXXCO

DOCUMENT TYPE:

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT      | PATENT NO.                                      |     |      |     |     | KIND DATE |          |                 |     | APPL            | ICAT: | DATE |     |          |     |            |     |  |  |
|----------|---|-----|------|-----|-----|-----------|----------|-----------------|-----|-----------------|-------|------|-----|----------|-----|------------|-----|--|--|
|          |   |     |      |     | A1  | -         | 2006     | 0720            | 1   | US 2            | 005-3 | 3128 | 73  |          | 2   | 0051       | 220 |  |  |
| WO       | US 2006160748<br>WO 2005051978<br>WO 2005051978 |     |      | A2  |     | 20050609  |          | WO 2004-US39739 |     |                 |       |      |     | 20041126 |     |            |     |  |  |
| WO       |   |     |      |     | A3  |           | 20051124 |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          | W:  |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          | RW:   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     | BF,       | ВJ,      | CF,             | CG, | CI,             | CM,   | GΑ,  | GN, | GQ,      | GW, | ML,        | MR, |  |  |
|          |   | •   | •    | •   |     |           | -        | 14 81           | •   | -               |       |      |     | +        |     |            | -   |  |  |
| WO       | 20061<br>20050<br>W:<br>RW:<br>20070<br>W:      |     |      |     |     |           | 2007     |                 |     |                 |       |      |     |          |     | 00612      |     |  |  |
|          | ₩:  |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     | •         |          |                 | •   | •               | •     | SY,  | TJ, | TM,      | TN, | TR,        | TT, |  |  |
|          |   |     |      |     |     |           | •        |                 | •   |                 |       |      |     |          |     |            |     |  |  |
|          | RW:   |     |      |     |     |           |          |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           |          | -               | -   |                 |       |      | •   | •        | •   |            |     |  |  |
|          |   |     |      |     |     |           | GN,      |                 |     |                 |       |      |     |          |     |            |     |  |  |
|          |   |     |      |     |     |           | NA,      | SD,             | SL, | SZ,             | TZ,   | UG,  | ZM, | ZW,      | AM, | ΑŻ,        | BY, |  |  |
| DD T     |   | -   |      | MD, | RU, | TJ,       | TM       |                 | _   |                 |       |      |     |          |     |            |     |  |  |
| PRIORITY | Y APP   | LN. | INFO | . : |     |           |          |                 | ]   | US 2003-524833P |       |      |     |          |     | P 20031125 |     |  |  |

WO 2004-US39739 A2 20041126 AB The invention discloses compds. containing single amino acids, peptides or their derivs. which are are selectively delivered to the mitochondria of a cell. These compds. exhibit antioxidant activity thereby reducing reactive oxygen species in cells and are useful for inhibiting oxidative stress-induced cell injury or death both in vivo and ex vivo. Thus, N-acetyl-L-cysteine choline ester was prepared and shown to minimize the depolarization of mitochondrial membrane potential induced by oxidative stress.

IT 853015-41-5P 853015-46-0P 853015-49-3P
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
 (Uses)

(preparation of compds. for delivering amino acids or peptides with antioxidant activity into mitochondria)

RN 853015-41-5 CAPLUS

CN Glycine, L-γ-glutamyl-L-cysteinyl-, 3-[2-(trimethylammonio)ethyl]
 ester, chloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$Me_3+N$$
 $O$ 
 $N$ 
 $H$ 
 $R$ 
 $H$ 
 $S$ 
 $CO_2H$ 

• c1-

RN 853015-46-0 CAPLUS

CN Ethanaminium, 2-[(2R)-2-(acetylamino)-3-mercapto-1-oxopropoxy]-N,N,N-trimethyl-, chloride (1:1) (CA INDEX NAME)

Absolute stereochemistry.

● cl -

RN 853015-49-3 CAPLUS

CN Glycine, L-γ-glutamyl-L-cysteinyl-, 3-[2-(trimethylammonio)ethyl]
 ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

ED Entered STN: 09 Jun 2006

ACCESSION NUMBER: 2006:544676 CAPLUS

DOCUMENT NUMBER:

145:21218

TITLE:

Compositions and methods for attenuating mitochondria-mediated cell injury comprising

S-nitrosated thiol antioxidants

INVENTOR(S):

Brookes, Paul S.; Sheu, Shey-Shing; Anders, Marion W.

PATENT ASSIGNEE(S):

University of Rochester, USA

SOURCE:

U.S. Pat. Appl. Publ., 12 pp., Cont.-in-part of

PCT/US04/39739.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| PA      | TENT     | NO.                      |      |       | KIND DATE |     |                      |     | APPL  | ICAT                              |                | DATE      |          |     |          |       |     |  |  |
|---------|----------|--------------------------|------|-------|-----------|-----|----------------------|-----|-------|-----------------------------------|----------------|-----------|----------|-----|----------|-------|-----|--|--|
| US      | 2006     | 2006122267               |      |       |           |     | A1 2006              |     |       | US 2                              | <br>005-316618 |           |          |     | 20051220 |       |     |  |  |
| WO      | 2005     | 2005051978<br>2005051978 |      |       | A2        |     | 20050609<br>20051124 |     |       | WO 2                              | 004-1          |           | 20041126 |     |          |       |     |  |  |
| WO      | 2005     |                          |      |       | A3        |     |                      |     |       |                                   |                |           |          |     |          |       |     |  |  |
|         | W:       | ΑE,                      | AG,  | ΑL,   | AM,       | ΑT, | ΑU,                  | ΑZ, | BA,   | BB,                               | BG,            | BR,       | BW,      | BY, | BZ,      | CA,   | CH, |  |  |
|         |          | CN,                      | CO,  | CR,   | CU,       | CZ, | DE,                  | DK, | DM,   | DZ,                               | EC,            | EE,       | EG,      | ES, | FI,      | GB,   | GD, |  |  |
|         |          | GE,                      | GH,  | GM,   | HR,       | HU, | ID,                  | IL, | IN,   | IS,                               | JP,            | KE,       | KG,      | KP, | KR,      | KZ,   | LC, |  |  |
|         |          | LK,                      | LR,  | LS,   | LT,       | LU, | LV,                  | MA, | MD,   | MG,                               | MK,            | MN,       | MW,      | MX, | MZ,      | NA,   | NI, |  |  |
|         |          |                          |      |       |           |     | PL,                  |     |       |                                   |                |           |          |     |          |       |     |  |  |
|         |          | ТJ,                      | TM,  | TN,   | TR,       | TT, | ΤZ,                  | ŪΑ, | UG,   | US,                               | UΖ,            | VC,       | VN,      | YU, | ZA,      | ZM,   | ZW  |  |  |
|         | RW:      | BW,                      | GH,  | GM,   | KΕ,       | LS, | MW,                  | ΜZ, | NA,   | SD,                               | SL,            | SZ,       | TZ,      | UG, | ZM,      | ZW,   | AM, |  |  |
|         |          | ΑZ,                      | BY,  | KG,   | KZ,       | MD, | RU,                  | ТJ, | TM,   | ΑT,                               | BE,            | BG,       | CH,      | CY, | CZ,      | DE,   | DK, |  |  |
|         |          | EE,                      | ES,  | FI,   | FR,       | GB, | GR,                  | HU, | ΙE,   | IS,                               | IT,            | LU,       | MC,      | NL, | PL,      | PT,   | RO, |  |  |
|         |          | SE,                      | SI,  | SK,   | TR,       | BF, | ВJ,                  | CF, | CG,   | CI,                               | CM,            | GΑ,       | GN,      | GQ, | GW,      | ML,   | MR, |  |  |
|         |          | ΝE,                      | SN,  | TD,   | TG        |     |                      |     |       |                                   |                |           |          | ,   |          | •     | -   |  |  |
| MO      | 2007     |                          |      |       | A2        |     | 2007                 |     |       |                                   |                |           |          |     |          | 0061  |     |  |  |
|         | W :      | ΑE,                      | AG,  | AL,   | AM,       | ΑT, | ΑU,                  | AZ, | BA,   | BB,                               | BG,            | BR,       | BW,      | BY, | ΒZ,      | CA,   | CH, |  |  |
|         |          | CN,                      | CO,  | CR,   | CU,       | CZ, | DE,                  | DK, | DM,   | DZ,                               | EC,            | EE,       | EG,      | ES, | FI,      | GB,   | GD, |  |  |
|         |          | GE,                      | GH,  | GM,   | GT,       | HN, | HR,                  | ΗU, | ID,   | IL,                               | IN,            | IS,       | JP,      | KE, | KG,      | KM,   | KN, |  |  |
|         |          | ΚP,                      | KR,  | KZ,   | LA,       | LC, | LK,                  | LR, | LS,   | LT,                               | LU,            | LV,       | LY,      | MA, | MD,      | MG,   | MK, |  |  |
|         |          | -MN ;-                   | MW,  | -MX-, | MY;       | MZ; | -NA,                 | NG, | ~N·I, | NO,                               | ··NZ;          | OM,       | PG,      | PH, | PL,      | PT,   | RO, |  |  |
|         |          | RS,                      | RU,  | SC,   | SD,       | SE, | SG,                  | SK, | SL,   | SM,                               | sv,            | SY,       | TJ,      | TM, | TN,      | TR,   | TT, |  |  |
|         |          |                          |      |       |           |     | VC,                  |     |       |                                   |                |           |          |     |          |       |     |  |  |
|         | RW:      | AT,                      | BE,  | BG,   | CH,       | CY, | CZ,                  | DE, | DK,   | EE,                               | ES,            | FI,       | FR,      | GB, | GR,      | ΗU,   | ΙE, |  |  |
|         |          | IS,                      | IT,  | LT,   | LU,       | LV, | MC,                  | NL, | PL,   | PT,                               | RO,            | SE,       | SI,      | SK, | TR,      | BF,   | ВJ, |  |  |
|         |          |                          |      |       |           |     | GN,                  |     |       |                                   |                |           |          |     |          |       |     |  |  |
|         |          | GM,                      | KE,  | LS,   | MW,       | MZ, | NA,                  | SD, | SL,   | SZ,                               | TZ,            | ŪĠ,       | ZM,      | ZW, | AM,      | ΑZ,   | BY, |  |  |
| DDTODIM | ים מול ע |                          |      | MD,   | Rυ,       | TJ, | TM                   |     |       |                                   |                |           |          |     |          |       |     |  |  |
| PRIORIT | I APP.   | ⊔Ν                       | TMLO | . :   |           |     |                      |     |       | US 20                             |                |           |          | ]   |          |       |     |  |  |
|         |          |                          |      |       |           |     |                      |     |       | WO 2004-US39739<br>US 2005-316618 |                |           |          |     |          | 0041  |     |  |  |
| 3.D =1  |          |                          |      |       |           |     |                      |     | ,     | US 20                             | 105-3          | 3 T 6 6 : | r g      | 1   | A 20     | 00512 | 220 |  |  |

AB The present invention relates to an S-nitrosated mitochondria-targeted thiol-based antioxidant prodrug and uses therefore for the prevention or treatment of diseases or conditions associated with mitochondrial dysfunction resulting from changes in the mitochondrial redox environment. When activated, prodrug of the present invention can specifically provide a

NO• donor and a thiol-based antioxidant to mitochondria thereby decreasing the degree of mitochondrial dysfunction. Thus, a mitochondria-targeted NO. donor, S-nitroso-2-mercaptopropylglycine (SNO-MPG), obtained by S-nitrosation of 2-mercaptopropylglycine, protected cardiomyocytes from ischemia-reperfusion injury in a dose-dependent manner, with the higher dose being more effective than ischemic preconditioning (IPC).

IT 853177-82-9

RL: RCT (Reactant); RACT (Reactant or reagent)

(S-nitrosated mitochondria-targeted thiol antioxidant prodrugs for attenuating mitochondria-mediated cell injury as nitric oxide donors)

RN 853177-82-9 CAPLUS

CN Ethanaminium, 2-[(2R)-2-(acetylamino)-3-mercapto-1-oxopropoxy]-N,N,N-trimethyl- (9CI) (CA INDEX NAME)

## Absolute stereochemistry.

L4 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

ED Entered STN: 09 Jun 2005

ACCESSION NUMBER: 2005:490376 CAPLUS

DOCUMENT NUMBER:

143:26888

TITLE:

Preparation of compounds for delivering amino acids or peptides with antioxidant activity into mitochondria

INVENTOR(S):

Sheu, Shey-Shing; Anders, Marion W.; Xu, Lin; Sharma,

Virendra K.

PATENT ASSIGNEE(S):

University of Rochester, USA

SOURCE:

PCT Int. Appl., 51 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT    | CENT 1        | NO. |     |             | KIND DATE |     |      |      | i  | APPL  | ICAT:         |     | DATE         |       |          |     |     |  |
|--------|---------------|-----|-----|-------------|-----------|-----|------|------|--|-------|---------------|-----|--------------|-------|----------|-----|-----|--|
| <br>WO | O 2005051978  |     |     |             | A2        |     |      | 0609 | ,  | 7O 20 | <br>0 0 4 - 1 |     | 20041126     |       |          |     |     |  |
| WO     | NO 2005051978 |     |     | A3 20051124 |           |     |      |      |  |       |               |     |              |       |          |     |     |  |
|        | W;            | ΑE, | AG, | AL,         | AM,       | AT, | ΑU,  | ΑZ,  | BA,                                      | BB,   | BG,           | BR, | BW,          | BY,   | ΒZ,      | CA, | CH, |  |
|        |               |     |     |             |           |     | DE,  |      |  |       |               |     |              |       |          |     |     |  |
| <br>   |               | GE, | GH, | GM,-        | HR,       | HU, | ID,  | IL.  | IN.                                      | IS.   | JP;           | KE. | KG.          | KP.   | KR-      | KZ. | LC, |  |
|        |               |     |     |             |           |     | LV,  |      |  |       |               |     |              |       |          | -   |     |  |
|        |               | -   | -   |             | -         |     | PL,  | -    |  | •     |               |     | •            |       |          | •   | •   |  |
|        |               |     |     |             |           |     | TZ,  |      |  |       |               |     |              |       |          |     |     |  |
|        | RW:           | BW, |     |             |           |     |      |      |  |       |               |     |              | -     | -        | •   |     |  |
|        | ,             |     |     |             |           |     | RU,  |      |  |       |               |     |              |       |          |     |     |  |
|        |               |     |     |             |           |     | GR,  |      |  |       |               |     |              |       |          |     |     |  |
|        |               |     |     |             |           |     | ВJ,  |      |  |       |               |     |              |       |          |     |     |  |
|        |               |     |     | TD,         |           | ,   | _ ,  | ,    | ,  | ,     | ,             | ,   | ,            | - = / | •,       | ,   | /   |  |
| ΑU     | 2004          |     |     | •           |           |     | 2005 | 0609 | 7  | AU 20 | 004-          |     | 20041126     |       |          |     |     |  |
|        | CA 2547086    |     |     |             |           |     | 2005 |      |  |       |               |     | <del>-</del> |       |          |     |     |  |
| ΕP     | EP 1691818    |     |     |             | A2        |     |      |      | EP 2004-812293                           |       |               |     |              |       |          |     |     |  |
|        | R:            | AT, | BE, | CH,         | DE.       |     |      |      |  |       |               |     |              |       |          |     |     |  |
|        |               |     |     |             |           |     |      |      |  |       |               |     |              |       | ,        | ,   | ,   |  |
| US     | 2006          |     |     |             |           |     |      |      | CZ, EE, HU, PL, SK, IS<br>US 2005-316618 |       |               |     |              |       | 20051220 |     |     |  |
|        |               |     |     |             |           |     |      |      | US 2005-312873                           |       |               |     |              |       |          |     |     |  |
|        |               |     |     |             |           |     |      |      |  |       |               |     |              |       | ~ .      |     |     |  |

US 2007099845 PRIORITY APPLN. INFO.:

A1 20070503

US 2006-580803

20061124

US 2003-524833P WO 2004-US39739 P 20031125 W 20041126

OTHER SOURCE(S): CASREACT 143:26888; MARPAT 143:26888

The invention discloses compds. containing single amino acids, peptides or their derivs. which have the potential to express antioxidant activity capable of reducing reactive oxygen species in cells. Compds.

R-O-Z-N+Q1Q2Q3 [R is an amino acid or a peptide comprising two or more amino acids (or their derivs.) which have antioxidant activity; Z is a linker mol. containing 1-20 atoms in a direct chain; Q1, Q2 and Q3 are independently aliphatic C1-C5 hydrocarbons; or Q2 is optional or Q2 and Q3 together form an aliphatic N-heterocycle] and related cyclic compds. containing N+Q1Q2 and having R-O-Z- as substituent. These compds. may be used to inhibit oxidative stress-induced cell injury or death both in vivo and ex vivo. Thus, N-acetyl-L-cysteine choline ester was prepared and shown to prevent the depolarization of membrane potential in isolated heart mitochondria induced by rotenone- and tert-Bu hydroperoxide-induced oxidative stress.

IT 853015-41-5P 853015-46-0P 853015-49-3P 853015-50-6P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of compds. for delivering amino acids or peptides with antioxidant activity into mitochondria)

RN 853015-41-5 CAPLUS

CN Glycine, L- $\gamma$ -glutamyl-L-cysteinyl-, 3-[2-(trimethylammonio)ethyl] ester, chloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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RN 853015-46-0 CAPLUS

CN Ethanaminium, 2-[(2R)-2-(acetylamino)-3-mercapto-1-oxopropoxy]-N,N,N-trimethyl-, chloride (1:1) (CA INDEX NAME)

Absolute stereochemistry.

● cl -

RN 853015-49-3 CAPLUS

CN Glycine,  $L-\gamma$ -glutamyl-L-cysteinyl-, 3-[2-(trimethylammonio)ethyl]